

RESEARCH

Open Access



The impact of gestational weight gain on fetal and neonatal outcomes: the Araraquara Cohort Study

Audêncio Victor^{1,2,4*} , Laísia de França da Silva Teles², Isabel Oliveira Aires², Leticia Falcão de Carvalho², Liania A. Luzia², Rinaldo Artes³ and Patrícia H. Rondó^{1,2}

Abstract

Background Gestational weight gain (GWG) is an important indicator for monitoring maternal and fetal health. Objective: To evaluate the effect of GWG outside the recommendations of the Institute of Medicine (IOM) on fetal and neonatal outcomes.

Study design A prospective cohort study with 1642 pregnant women selected from 2017 to 2023, with gestational age \leq 18 weeks and followed until delivery in the city of Araraquara, Southeast Brazil. The relationship between IOM-recommended GWG and fetal outcomes (abdominal subcutaneous tissue thickness, arm and thigh subcutaneous tissue area and intrauterine growth restriction) and neonatal outcomes (percentage of fat mass, fat-free mass, birth weight and length, ponderal index, weight adequateness for gestational age by the Intergrowth curve, prematurity, and Apgar score) were investigated. Generalized Estimating Equations were used.

Results GWG below the IOM recommendations was associated with increased risks of intrauterine growth restriction (IUGR) (aOR 1.61; 95% CI: 1.14–2.27), low birth weight (aOR 2.44; 95% CI: 1.85–3.21), and prematurity (aOR 2.35; 95% CI: 1.81–3.05), and lower chance of being Large for Gestational Age (LGA) (aOR 0.38; 95% CI: 0.28–0.54), with smaller arm subcutaneous tissue area (AST) (-7.99 g; 95% CI: -8.97 to -7.02), birth length (-0.76 cm; 95% CI: -1.03 to -0.49), and neonatal fat mass percentage (-0.85%; 95% CI: -1.12 to -0.58). Conversely, exceeding GWG guidelines increased the likelihood of LGA (aOR 1.53; 95% CI: 1.20–1.96), with lower 5th-minute Apgar score (aOR 0.42; 95% CI: 0.20–0.87), and increased birth weight (90.14 g; 95% CI: 53.30 to 126.99).

Conclusion Adherence to GWG recommendations is crucial, with deviations negatively impacting fetal health. Effective weight control strategies are imperative.

Keywords Gestational weight gain, Pregnancy, Fetal outcomes, Neonatal outcomes, Cohort study, Intrauterine growth restriction

*Correspondence:

Audêncio Victor
Audenciovictor@gmail.com

¹Public Health Postgraduate Program, School of Public Health, University of São Paulo, São Paulo, SP, Brazil

²Nutrition Department, School of Public Health, University of São Paulo, São Paulo, SP, Brazil

³Inspier - Institute of Education and Research, São Paulo, Brazil

⁴Faculdade de Saúde Pública- USP, Avenida Doutor Arnaldo, 715 – São Paulo, São Paulo, Brazil



© The Author(s) 2024. **Open Access** This article is licensed under a Creative Commons Attribution 4.0 International License, which permits use, sharing, adaptation, distribution and reproduction in any medium or format, as long as you give appropriate credit to the original author(s) and the source, provide a link to the Creative Commons licence, and indicate if changes were made. The images or other third party material in this article are included in the article's Creative Commons licence, unless indicated otherwise in a credit line to the material. If material is not included in the article's Creative Commons licence and your intended use is not permitted by statutory regulation or exceeds the permitted use, you will need to obtain permission directly from the copyright holder. To view a copy of this licence, visit <http://creativecommons.org/licenses/by/4.0/>. The Creative Commons Public Domain Dedication waiver (<http://creativecommons.org/publicdomain/zero/1.0/>) applies to the data made available in this article, unless otherwise stated in a credit line to the data.

Introduction

Pregnancy is a critical window in maternal and child health, where gestational weight gain (GWG) emerges as a determinant factor for fetal well-being and development. Monitoring GWG is essential, serving as a prognostic marker for the health of the pregnant woman and the conceptus [1–5].

Guidelines for GWG have changed over time, reflecting the understanding of its importance. In the 1950s, the recommendation was for a gain of 10–14 lb, aiming to prevent complications such as pre-eclampsia (Ferguson et al., 1950). The following decade recognized the positive association between higher GWG and birth weight [5]. In 1990, the Institute of Medicine (IOM) of the USA established specific recommendations, with the most recent update in 2019, emphasizing the importance of not exceeding GWG targets to avoid obesity-related complications [6]. To mitigate adverse impacts on maternal and fetal health, the IOM in 2009 outlined guidelines suggesting a total weight gain of 11 to 16 kg for women of normal weight, 7 to 11 kg for overweight women, and 5 to 9 kg for obese women [7].

Excessive or insufficient GWG can increase the risk of negative outcomes for both the mother and the conceptus [8–10]. Research indicates that GWG above IOM recommendations is associated with an increased risk of metabolic complications, hypertension, gestational diabetes, cesarean section, postpartum weight retention, macrosomia, childhood obesity, and cardiometabolic outcomes in childhood [8–10]. Conversely, GWG below the recommendations is linked to an elevated risk of intrauterine growth restriction (IUGR), low birth weight (LBW), preterm birth, perinatal mortality, and a higher incidence of newborns small for gestational age (SGA) [11–13]. Furthermore, studies indicate that GWG within the IOM-recommended ranges is associated with a lower incidence of LBW and better health outcomes [14].

The possibility of mitigating the risks of gestational complications and adverse fetal outcomes in obese pregnant women by limiting GWG is an area of growing interest [15, 16]. The literature has highlighted the importance of adjusting GWG in various interventions, including diet, physical activity, the use of probiotics, and psychological and behavioral approaches [17].

In Brazil, although there are no standardized recommendations for GWG based on Brazilian population data, new guidelines for GWG have been proposed based on national GWG curves. Thus, the IOM recommendations, widely used to classify GWG, still constitute a viable alternative. Given this context, the objective of this study is to evaluate whether pregnant women with GWG outside the IOM-recommended range present adverse fetal and neonatal outcomes compared to those whose GWG is considered adequate, aiming to provide support

for clinical practice and contribute to the development of effective intervention strategies that promote a healthy pregnancy and minimize risks to the mother and respective conceptus.

Materials and methods

Study design and participants

A population-based prospective epidemiological cohort study, part of the ongoing “Araraquara Cohort Study”, was conducted. The sample included pregnant women with gestational age ≤ 18 weeks, recruited from Health Units in Araraquara, São Paulo, Brazil, and monitored quarterly until delivery from 2017 to 2023. Women with multiple pregnancies or who experienced miscarriage were excluded. Pregnant women lacking information on height, pre-pregnancy weight, and weight at delivery were also excluded (Fig. 1).

Main exposure: GWG

GWG was calculated from the difference between weight at delivery and pre-pregnancy weight, with classifications according to IOM recommendations. Pre-pregnancy BMI was determined from measurements taken at the first assessment. For women with gestational age < 13 weeks, current weight was used; for those ≥ 13 weeks, weight recorded at the first prenatal visit was used. Nutritional status classification followed WHO guidelines: underweight (BMI < 18.5 kg/m²), normal weight (18.5–24.9 kg/m²), overweight (25–29.9 kg/m²), or obesity (≥ 30 kg/m²) [18].

Fetal outcomes

Fetal variables were assessed longitudinally throughout the three trimesters of pregnancy, including abdominal subcutaneous tissue thickness (AST), arm subcutaneous tissue area (ASA), thigh subcutaneous tissue area (TSA), estimated fetal weight (EFW), and the diagnosis of IUGR, the latter based on the comparison of EFW with the fetal curve of the Intergrowth-21st Project. The diagnosis of IUGR was considered when the EFW was below the 10th percentile of the fetal curve [19, 20].

Neonatal outcomes

The neonatal outcomes evaluated were the percentage of fat mass (%FM), percentage of fat-free mass (%FFM), LBW, birth weight (g), birth length (cm), ponderal index (length/weight³), gestational age at birth (preterm and term), and adequateness of birth weight according to the Intergrowth-21st - large for gestational age (LGA) (> 90 th percentile), SGA (< 10 th percentile), and adequate for gestational age (AGA) (between the 10th and 90th percentiles) [20]. The body composition of the neonates, including %FM and %FFM, was assessed using air displacement plethysmography.

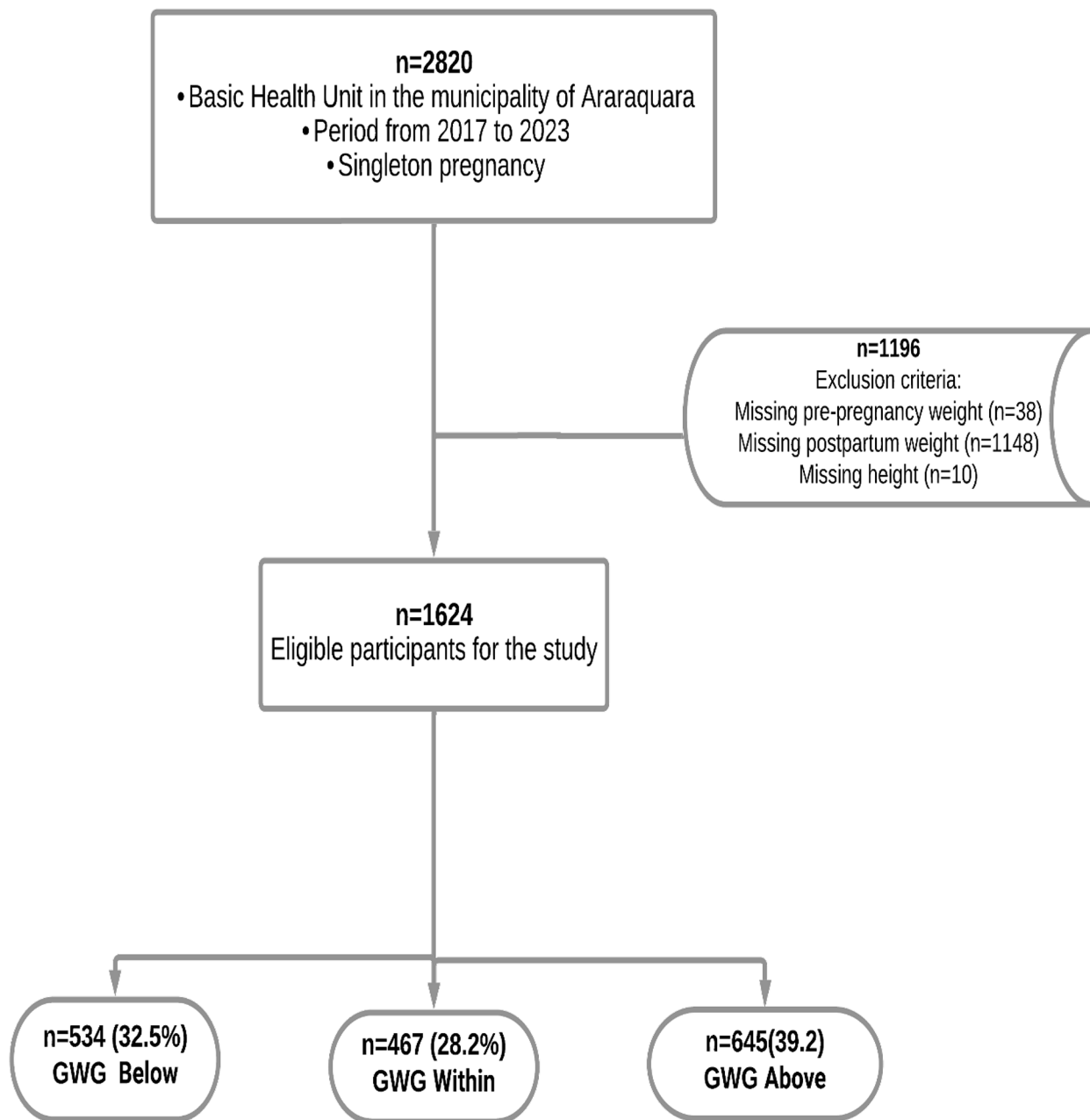


Fig. 1 Selection of study population in a cohort study in Araraquara

Maternal characteristics (study covariates)

Various factors were considered as covariates, including socioeconomic and demographic characteristics: age (years), education (years), number of people per room, number of previous delivery, parity, per capita income in reais (1 US\$ = 4.9 R\$), race (white or non-white), and marital status (married/in a stable union, single/separated/widowed). Lifestyle habits such as physical activity, smoking, and alcohol consumption were also considered, as well as morbidity (diabetes, hypertension, urinary tract

infection). The anthropometry of pregnant women (pre-pregnancy weight, current BMI), gestational age, glycemic profile (fasting glucose, insulin, HOMA, glyated hemoglobin), high-sensitivity C-reactive protein (hs-CRP), hemoglobin, and lipid profile (triglycerides, total cholesterol, and fractions).

Statistical analysis

A directed acyclic graph (DAG) was created to visually represent the theoretical model and clarify the

relationships between variables related to GWG, as well as fetal and neonatal outcomes. (Figure S1). The DAG was constructed using DAGitty software (version 3.0; Nijmegen, GE, Netherlands) [21]. Graphical criteria were applied for the selection of adjusted covariates, aiming to minimize bias in variable selection [22, 23].

Descriptive statistics was used for data analysis, and the Shapiro-Wilk test was utilized to check the normality of continuous variables. Non-normal continuous variables were presented as median and interquartile range (IQR), and categorical ones, in frequencies and percentages (n (%)). Bivariate analyses were conducted to examine associations between independent and dependent variables. The Kruskal-Wallis test was used for continuous variables and the chi-square or Fisher's exact test for categorical ones. Generalized Linear Models were applied to associate GWG with neonatal outcomes (linear regression for continuous and logistic for binary). For fetal outcomes, longitudinal analysis with Generalized Estimating Equation (GEE) was used, considering the correlation between repeated observations of each pregnant woman [24–26]. Statistical analyses investigated associations between GWG and the outcomes of interest, considering the different GWG recommendations. The results were presented in measures of association (OR and Beta coefficient (β)), with statistical significance indicated by $P < 0.05$ and 95% confidence intervals. All analyses were performed using R software (version 4.1.0, R Foundation for Statistical Computing, Vienna, Austria).

Results

Maternal characteristics

Table 1. In this study, the association between maternal characteristics and GWG was evaluated in 1642 Brazilian women. The median age of the women was 27 years (IQR: 22.1–31.8), and the median years of education were 11 years (IQR: 10–11). The majority were in a stable relationship ($n=1440$, 87.7%), with 54.1% ($n=889$) identifying as non-white. The per capita income was R\$ 667 (IQR: 409–1000), and the average number of previous delivery was 1 (IQR: 1–2).

The median gestational BMI increased from 25.8 kg/m² (IQR: 22.2–30.2) in the first trimester to 28.8 kg/m² (IQR: 25.6–32.9) in the third trimester. Adequate physical activity was maintained by 11% of the women throughout all trimesters ($n=181$). Smoking decreased from 7.9% ($n=129$) in the first trimester to 4.8% ($n=79$) in the third, and alcohol consumption from 20.3% ($n=333$) to 6.0% ($n=99$). Gestational diabetes increased from 5.0% ($n=82$) to 9.7% ($n=159$), showing an association with GWG.

Fasting insulin levels rose from 7.0 uIU/mL (IQR: 5–11) to 10 uIU/mL (IQR: 7–14) throughout pregnancy. Total cholesterol increased from 173 mg/dL

(IQR: 151–197) to 226 mg/dL (IQR: 195–258), LDL-c from 95.0 mg/dL (IQR: 77–113) to 127 mg/dL (IQR: 101–156), and triglycerides from 104 mg/dL (IQR: 81–134) to 186 mg/dL (IQR: 147–230), all associated with GWG. HDL-c varied from 56 mg/dL (IQR: 48–64.0) in the first period to 58 mg/dL (IQR: 50–68) in the third.

Neonatal and fetal characteristics

As shown in Table 2, fetal and neonatal outcomes were assessed in relation to GWG over the three periods of gestation. There was an increase in the AST, from 2.8 mm (IQR: 2.5–3.1) in the first period to 3.8 mm (IQR: 3.4–4.4) in the second trimester. The AST also showed growth, from 3.0 mm² (IQR: 2.4–3.6) to 6.6 mm² (IQR: 5.7–7.8), as did the TST, which increased from 5.4 mm² (IQR: 4.4–6.7) to 13.5 mm² (IQR: 11.4–15.9). The EFW increased from 171 g (IQR: 145–212) in the first period to 616 g (IQR: 486–776) in the second, reaching 2064 g (IQR: 1814–2341) in the third trimester. The prevalence of IUGR decreased from 16.3% ($n=268$) to 1.5% ($n=23$) from the first to the second period.

In neonatal outcomes, the %FM at birth was 21% (IQR: 17–24), and the %FFM was 79% (IQR: 76–83). The prevalence of prematurity was 9.3% ($n=154$), while most neonates were born at term (89.7%, $n=1474$). The Apgar score at the 5th minute was satisfactory (≥ 7) in 97.1% of cases ($n=1594$). The length at birth was 49.0 cm (IQR: 47–50), and the birth weight was 3222 g (IQR: 2895–3520). The adequateness of birth weight, according to the Intergrowth curve, showed 70.3% ($n=1155$) as AGA, 8.8% ($n=144$) as SGA, and 8.3% ($n=137$) as LGA. The incidence of LBW was 8.6% ($n=142$), with most neonates presenting with an adequate weight 90.6% ($n=1488$).

Association between GWG and fetal and neonatal outcomes

Tables 3 and 4 present the adjusted associations between GWG and fetal and neonatal outcomes. The results indicate that a GWG below the recommended range was associated with increased risks of IUGR (aOR 1.61; 95% CI: 1.14–2.27), LBW (aOR 2.44; 95% CI: 1.85–3.21), and prematurity (aOR 2.35; 95% CI: 1.81–3.05), and a lower chance of being LGA (aOR 0.38; 95% CI: 0.28–0.54), AST (-7.99 g; 95% CI: -8.97 to -7.02), birth length (-0.76 cm; 95% CI: -1.03 to -0.49), and neonatal fat mass percentage (-0.85%; 95% CI: -1.12 to -0.58). Whereas a GWG above the recommended range resulted in a higher likelihood of neonates being LGA (aOR 1.53; 95% CI: 1.20–1.96), with a lower Apgar score at the 5th minute (aOR 0.42; 95%

Table 1 Maternal characteristics associated with GWG in three trimesters and its relationship with IOM recommendations

| Charateristics | 1st trimester | 2st trimestre | 3rd trimestre |
|---|------------------|------------------|------------------|
| Age (years) | 27 (22.1–31.8) | | |
| Maternal Education (years) | 11(10–11) | | |
| Number of Previous delivery † | 1 (1–2) | | |
| Number of people per room | 0.6 (0.5–0.8) | | |
| Per Capita Income in Brazilian Real (R\$) † | 667 (409–1000) | | |
| Race | | | |
| White | 753 (45.9) | | |
| Non-white | 889 (54.1) | | |
| Marital Status, n(%) | | | |
| Married or in a stable relationship | 1440 (87.7) | | |
| Single, separated, or widowed | 202 (12.3) | | |
| Pre-gestational BMI (kg /m ²) † | 25.7 (22.2–30.4) | | |
| Gestational BMI (kg /m ²) † | 25.8 (22.2–30.2) | 27.4 (24.0–31.5) | 28.8 (25.6–32.9) |
| Gestational Age (weeks) † | 13.6 (12.3–15.3) | 23.6 (22.1–25.1) | 33(31.7–34.1) |
| Physical Activity | | | |
| Adequate | 181 (11) | 143 1 (8.7) | 181 (11) |
| Inadequate | 578 (35.2) | 578 (35.2) | 578 (35.2) |
| Smoking † | | | |
| No | 1513 (92.1) | 1422 (86.6) | 1308 (79.7) |
| Yes | 129 (7.0) | 97 (5.9) | 79 (4.8) |
| Alcohol Consumption | | | |
| No | 1309 (79.7) | 1384 (84.3) | 1288 (78.4) |
| Yes | 333 (20.3) | 135 (8.2)) | 99 (6.0) |
| Diabetes, n(%) † | | | |
| No | 1560 (95.0) | 1395 (85.0) | 1233 (75.1) |
| Yes | 82 (5.0) | 123 (7.5) | 159 (9.7) |
| Hypertension | | | |
| No | 1526 (92.9) | 1405 (85.6) | 1281 (78.0) |
| Yes | 116 (7.1) | 113 (6.9) | 111 (6.8) |
| Urinary Tract Infection | | | |
| No | 1449 (88.2) | 1414 (86.1) | 1306 (79.5) |
| Yes | 193 (11.8) | 104 (6.3) | 86 (5.2) |
| Cervicitis/Vaginitis | | | |
| No | 1531 (93.2) | 1424 (86.7) | 1322 (80.5) |
| Yes | 111 (6.8) | 94 (5.7) | 70 (4.3) |
| hs-CRP (ng/mL) | 6.0 (3.1–11.6) | 6.1 (3.4–11.6) † | 5 (3–10) † |
| Hemoglobin (g/dL) † | 12.5 (11.9–13.1) | 11.8 (11.3–12.4) | 11.8 (11.3–12.4) |
| Glycated Hemoglobin % | 5.1 (4.9–5.3) | 4.9 (4.6–5.1) | 5 (5–5) |
| Fasting Insulin (uUI/mL) † | 7.0 (5–11) | 8.0 (6–12) | 10 (7–14) |
| Cholesterol (mg/dL) | 173 (151–197) | 210 (184–239) | 226 (195–258) |
| HDL-c (mg/dL) | 56.0 (48–64.0) † | 61.0 (53–71) † | 58 (50–68) |
| LDL-c (mg/dL) | 95.0 (77–113) | 116.0 (95.5–139) | 127 (101–156) |
| Triglycerides (mg/dL) | 104 (81–134) | 146 (113–185) | 186 (147–230) † |

The data is presented as number (percentage) and median with interquartile range (25th percentile – 75th percentile)

† Statistical differences observed between gestational weight gain groups, tested with: Kruskal-Wallis test for continuous variables and χ^2 test, Fisher's test for categorical variables

Abbreviations: BMI: body mass index; LDL-c: low-density lipoprotein cholesterol; HDL-c: high-density lipoprotein cholesterol. 1 Brazilian Real (R\$) equals approximately 4.9 US dollars (US\$)

Table 2 Fetal and neonatal outcomes in the three trimesters according to GWG

| Outcomes | 1st trimester | 2st trimester | 3rd trimester | |
|-------------------|---|---------------|------------------|------------------|
| Fetal | Abdominal subcutaneous tissue thickness (TSA) | 2.8 (2.5–3.1) | 3.8 (3.4–4.4) | |
| | Arm subcutaneous tissue area (TSB) | 3.0 (2.4–3.6) | 6.6 (5.7–7.8) | |
| | Thigh subcutaneous tissue area (TSC) | 5.4 (4.4–6.7) | 13.5 (11.4–15.9) | |
| | Estimated fetal weight in grams † | 171 (145–212) | 616 (486–776) | 2064 (1814–2341) |
| | Intrauterine growth restriction† | | | |
| | No | 918 (55.9) | 1346 (82.0) | |
| | Yes | 268 (16.3) | 23 (1.5) | |
| Neonatal (a) | % Fat mass (%FM) † | | 21 (17–24) | |
| | % Fat-free mass (%FFM) † | | 79 (76–83) | |
| | Prematurity † | | | |
| | | No | | 1474 (89.7) |
| | | Yes | | 154 (9.3) |
| | Apgar score at 5 minutes | | | |
| | | No | | 21 (1.3) |
| | | Yes | | 1594 (97.1) |
| | Birth length (cm) † | | | 49.0 (47–50) |
| | Birth weight (g) † | | | 3222 (2895–3520) |
| | Adequateness of birth weight (Intergrowth) † | | | |
| | | SGA | | 1155 (70.3) |
| | | AGA | | 144 (8.8) |
| | | LGA | | 137 (8.3) |
| Low birth weight† | | | | |
| | No | | 1488 (90.6) | |
| | Yes | | 142 (8.6) | |

The data is presented as number (percentage) and median with interquartile range (25th percentile – 75th percentile)

† Statistical differences observed between gestational weight gain groups, tested with: Kruskal-Wallis test for continuous variables and χ^2 test, Fisher's test for categorical variables. (a) Variables measured at birth

Table 3 Unadjusted and adjusted analysis to assess adverse fetal and neonatal outcomes (Categorical) associated with GWG

| Outcomes | Gestacional weight gain (IOM-2019) | | | |
|---------------------------------------|------------------------------------|-------------------------|-------------------------|-------------------------|
| | Below | Above | Below | Above |
| | Crude OR (IC95%) | Crude OR (IC95%) | Adjusted OR (IC95%) | Adjusted OR (IC95%) |
| Intrauterine growth restriction (FGR) | 1.53 (1.11–2.10) | 1.27 (0.92–1.74) | 1.61 (1.14–2.27) | 1.26 (0.89–1.78) |
| Prematurity | 2.34 (1.81–3.02) | 1.19 (0.91–1.57) | 2.35 (1.81–3.05) | 1.24 (0.94–1.64) |
| Apgar score at 5 minutes | 0.58 (0.26–1.28) | 0.34 (0.16–0.71) | 0.52 (0.24–1.16) | 0.42 (0.20–0.87) |
| Small for gestational age (SGA) | 1.18 (0.92–1.50) | 0.78 (0.60–1.01) | 1.18 (0.92–1.53) | 0.77 (0.59–1.00) |
| Large for gestational age (LGA) | 0.43 (0.32–0.60) | 1.54 (1.21–1.95) | 0.38 (0.28–0.54) | 1.53 (1.20–1.96) |
| Low birth weight (LBW) | 2.32 (1.78–3.03) | 1.17 (0.88–1.55) | 2.44 (1.85–3.21) | 1.21 (0.91–1.62) |

The association between each fetal or neonatal outcome and GWG. Analyses adjusted for: Maternal age; marital status; race/color; mothers' education in years of study; BMI; gestational age in weeks; parity; number of rooms per person; lactose consumption; smoking; diabetes; serum levels of hemoglobin; glycated hemoglobin; cholesterol; triglycerides; HDL; LDL. Estimated odds ratios (OR) using a generalized estimating equation. OR, Odds Ratio; ORa, Adjusted Odds Ratio; CI, Confidence Interval

CI: 0.20–0.87), and a higher birth weight (90.14 g; 95% CI: 53.30 to 126.99).

Discussion

In this study, we found that GWG outside the IOM recommendations is significantly associated with adverse outcomes, both fetal and neonatal. It was observed that a GWG below the recommendations increases the risk of IUGR, LBW, and prematurity. These findings align with the hypothesis that inadequate nutrition during gestation can compromise fetal development, which is

corroborated by several studies [11–14]. On the other hand, a GWG above the recommendations was associated with an increased risk of LGA and changes in birth weight and length, suggesting that excess weight gain may predispose to obstetric complications and negatively impact the long-term health of the conceptus.

These findings corroborate previous studies that also found associations between GWG below and above the recommendations and adverse outcomes, indicating that inadequate GWG can have negative effects on the health of the fetus and newborn [14, 27–38].

Table 4 Unadjusted and adjusted analysis to assess adverse fetal and neonatal outcomes (Quantitative) associated with GWG

| Outcomes | Gestational weight gain (IOM-2019) | | | |
|---|------------------------------------|--------------------------|-------------------------------|-------------------------|
| | Below | Above | Below | Above |
| | β (IC95%) | β (IC95%) | β adj (IC95%) | β adj (IC95%) |
| Abdominal subcutaneous tissue thickness (TSA) | -0.02 (-0.10-0.06) | 0.01 (-0.06-0.09) | -0.00 (-0.06-0.06) | 0.05 (-0.00-0.11) |
| Arm subcutaneous tissue area (TSB) | -0.14 (-0.35-0.07) | 0.06 (-0.15-0.27) | -7.99 (-8.97-7.02) | -0.08 (-0.17-0.01) |
| Thigh subcutaneous tissue area (TSC) | -0.55 (-1.04 - -0.06) | -0.16 (-0.62-0.31) | -0.25 (-0.45-0.05) | -0.01 (-0.20-0.19) |
| % Fat mass (%FM) | -0.28 (-0.76-0.20) | 2.52 (2.09-2.95) | -0.85 (-1.12-0.58) | 1.21 (0.94-1.48) |
| % Fat-free mass (%FFM) | 1.51 (0.09-2.94) | 0.87 (-0.41-2.15) | 1.05 (-0.39-2.49) | 0.52 (-0.86-1.91) |
| Birth length | -0.79 (-1.07-0.51) | -0.22 (-0.56-0.12) | -0.76 (-1.03-0.49) | -0.26 (-0.60-0.08) |
| Fetal weight | -62.9 (-136.4-10.6) | -39.8 (-112.1-32.5) | -17.73 (-38.07-2.61) | 7.50 (-13.03-28.04) |
| Birth weight | -175.2 (-214.2-136.3) | 98.5 (61.5-135.4) | -189.30 (-227.9-150.7) | 90.14 (53.3-127) |

The association between each fetal or neonatal outcome and GWG. **Analyses adjusted for:** Maternal age; marital status; race/color; mothers' education in years of study; BMI; gestational age in weeks; parity; number of rooms per person; lactose consumption; smoking; diabetes; serum levels of hemoglobin; glycated hemoglobin; cholesterol; triglycerides; HDL; LDL. β , beta coefficient of the generalized estimating equation. CI, confidence interval

Similar to our results, Goldstein et al. [28] in a systematic review and meta-analysis that investigated the association between GWG, and maternal and infant outcomes observed that inadequate GWG, whether below or above the IOM recommendations, is associated with adverse outcomes. Similar findings were reported in other studies [33, 35, 39]. The issue of prematurity deserves special attention, as women with premature babies have less time to gain weight during pregnancy, which may indicate reverse causality in the association between GWG and prematurity.

Liu et al. [30], in a retrospective cohort study with a sample of 9 million mother-child pairs to investigate the associations between GWG and adverse birth outcomes, showed that both insufficient and excessive GWG were associated with a higher risk of adverse outcomes, such as LBW, prematurity, and SGA. These findings highlight the importance of achieving an adequate balance in GWG, avoiding both restriction and excessive weight gain during gestation.

Furthermore, the relationship between GWG and adverse outcomes may be influenced by other factors, such as pre-gestational BMI. Studies by Chiavaroli et al. [32] and Liu et al. [30] reported that both maternal BMI and GWG are associated with adverse outcomes, suggesting that weight control before and during gestation is crucial. These findings are supported by other research, such as the study by Athukorala et al. [33], which found an increased risk of adverse outcomes in pregnant women with overweight or obesity, including prematurity and LBW.

However, it is important to note that GWG is just one of many factors that can influence birth weight. Other factors, such as maternal health, nutrition, smoking, and alcohol consumption, can also play a significant role. Additionally, the effect of GWG may vary in different populations, depending on factors such as BMI, age, ethnicity, parity, and underlying medical conditions of the mother [40].

Similar to our results, Goldstein et al. (2017b) in a systematic review and meta-analysis that investigated the association between GWG and maternal and infant outcomes observed that inadequate GWG, whether below or above the IOM recommendations, is associated with adverse outcomes. Similar findings were reported in other studies (Athukorala et al., 2010b; Chowdhury et al., 2022; Truong et al., 2015b). The issue of prematurity deserves special attention, as women with premature babies have less time to gain weight during pregnancy, which may indicate reverse causality in the association between GWG and prematurity.

Liu et al. (2022), in a retrospective cohort study with a sample of 9 million mother-child pairs to investigate the associations between GWG and adverse birth outcomes, showed that both insufficient and excessive GWG were associated with a higher risk of adverse outcomes, such as LBW, prematurity, and SGA. These findings highlight the importance of achieving an adequate balance in GWG, avoiding both restriction and excessive weight gain during gestation.

Furthermore, the relationship between GWG and adverse outcomes may be influenced by other factors, such as pre-gestational BMI. Studies by Chiavaroli et al. (2021) and Liu et al. (2022) reported that both maternal BMI and GWG are associated with adverse outcomes, suggesting that weight control before and during gestation is crucial. These findings are supported by other research, such as the study by Athukorala et al. (2010b), which found an increased risk of adverse outcomes in pregnant women with overweight or obesity, including prematurity and LBW.

However, it is important to note that GWG is just one of many factors that can influence birth weight. Other factors, such as maternal health, nutrition, smoking, and alcohol consumption, can also play a significant role. Additionally, the effect of GWG may vary in different populations, depending on factors such as BMI, age,

ethnicity, parity, and underlying medical conditions of the mother [40],

In Brazil, a systematic review by Godoy et al. (2015) analyzed the recommendations for GWG and its impact on fetal outcomes. It was observed that both GWG below and above the recommendations are associated with adverse outcomes. Women with pre-gestational overweight or obesity have a higher risk of excessive gestational weight gain, which is associated with fetal macrosomia and high cesarean rates. Conversely, insufficient GWG can also lead to complications such as LBW and IUGR. In studies conducted in the Northeast and Southeast regions of Brazil, an association between macrosomia and women with pre-gestational overweight or obesity was observed [41–43]. In one of these studies, it was found that women with excessive GWG were more likely to have newborns with macrosomia [44]. Not only excessive weight and obesity influence perinatal outcomes, but insufficient GWG can also lead to complications, such as LBW and IUGR. Women with low pre-gestational body weight are more likely to have babies with LBW [45]. These findings highlight the importance of maintaining adequate GWG in Brazil to improve perinatal outcomes and reduce complications such as macrosomia and LBW. However, our study stands out by considering specific contextual variables of the Brazilian population, which may influence GWG and its outcomes. For example, the research by Drehmer et al. [46], in Brazil also noted the importance of the socioeconomic context in GWG and its effects on newborn health, but our study expands this understanding by including more detailed analyses of fetal and neonatal outcomes. Another point of distinction is our analysis of variables such as physical activity and smoking, which are known lifestyle factors that influence GWG. Studies like that of Nascimento et al. [47], in Brazil have shown that physical activity during pregnancy can help maintain GWG within the recommendations, which is consistent with our observations that maintaining adequate physical activity was low among women in the study.

Since GWG is a modifiable risk factor, it is possible to identify and correct its variations during gestation. Although various studies have demonstrated that appropriate interventions based on dietary changes can be effective in controlling weight gain and reducing the risk of complications during gestation [29], promoting healthy GWG can be an important element in avoiding adverse consequences for both the mother and the conceptus. Another important reference is the study by Dodd et al. [40], which conducted a systematic review of randomized clinical trials on antenatal interventions for overweight or obese pregnant women. The authors concluded that interventions such as dietary counseling and physical activity during gestation can result in modest reductions

in maternal weight gain but did not have a consistent effect on the perinatal outcomes assessed. The target of the intervention should be women with abnormal weight, more prone to having adverse outcomes in future pregnancies. It is also concerning the increased risks of preterm birth, LBW, and IUGR in pregnant women with low weight, compared to those of normal weight. Even in the normal weight group, the risk of cesarean section, preterm birth, and excessive fetal growth increases with the elevation of BMI. Therefore, it is important to be concerned with women with overweight or obesity before gestation, but it is also necessary to strengthen the management of women with low weight and normal weight, who are often neglected. The findings of this study reinforce the need for clear and personalized guidelines for GWG, as well as the importance of regular weight monitoring during gestation. Nutritional and lifestyle interventions should be prioritized to help pregnant women achieve a healthy GWG.

In this study, we identified a notable and unprecedented finding: a significant association between insufficient GWG and reduction in the area of the fetal subcutaneous tissue of the arm but no reduction in the area of the subcutaneous tissue of the thigh or the thickness of the abdominal subcutaneous tissue, suggesting a regionalized response of fetal adipose tissue to maternal nutrition. This phenomenon may have important implications for neonatal health, as adipose tissue plays an essential role in energy reserves and thermoregulation of the newborn [48]. The uneven distribution of fetal adipose tissue may reflect an adaptation to a restrictive intrauterine environment, which may predispose the child to long-term metabolic and health challenges [49]. Furthermore, metabolic programming, which can be influenced by GWG, is a potential mechanism that may explain variations in the accumulation of fetal adipose tissue and its consequences for child health [50]. Maternal obesity and excessive GWG have been associated with epigenetic changes that may affect fetal development and the risk of metabolic diseases later in life [51]. On the other hand, prenatal exposure to insufficient GWG and very severe maternal obesity have also been associated with adverse neuropsychiatric outcomes in children, highlighting the importance of nutritional balance during gestation [52, 53].

Limitations of this study include the lack of dietary data from pregnant women and the observational nature of the study, which prevents the inference of direct causal relationships. Additionally, despite adjustment for various covariates, we cannot rule out the possibility of residual bias. Future research should focus on better understanding the biological mechanisms linking GWG to adverse outcomes and on developing effective interventions to promote healthy GWG. Despite these

limitations, the study has potentialities, such as being prospective and population-based, which allows the evaluation of a wide range of maternal, fetal, and neonatal outcomes. The prospective cohort approach and the use of objective measures, such as ultrasonography and air displacement plethysmography, contribute to obtaining accurate information about fetal development. The inclusion of various covariates related to socioeconomic, demographic, lifestyle, obstetric history, and clinical profile of pregnant women allows a comprehensive analysis of the factors associated with the outcomes evaluated.

Therefore, the need for more personalized GWG guidelines and the importance of targeted nutritional interventions during gestation are reinforced by these findings. Understanding the underlying mechanisms governing the distribution of fetal adipose tissue and its implications for child development and long-term health is crucial for optimizing perinatal outcomes and for the formulation of effective public health policies.

Conclusion

This study underscores the imperative for targeted policies and interventions to manage GWG, which is critical for fetal and neonatal health. Effective weight monitoring and control strategies are essential to prevent insufficient or excessive gains, fostering a healthy pregnancy and averting neonatal complications. Access to information and support regarding appropriate GWG is crucial for maternal-infant well-being, and early intervention may forestall long-term health issues.

A novel finding of this investigation is the association between inadequate GWG and a reduction in the fetal subcutaneous arm tissue area, indicating a regionalized response of fetal adipose tissue to maternal nutrition. This could have significant implications for neonatal health and child development, suggesting that the distribution of fetal adipose tissue is a sensitive marker of the intrauterine environment. GWG guidelines should be tailored to reflect these insights, with a focus on personalized, evidence-based approaches. It is imperative that women achieve healthy weight targets during pregnancy, considering the regionalized response of fetal adipose tissue in the development of future nutritional guidelines and interventions.

Supplementary Information

The online version contains supplementary material available at <https://doi.org/10.1186/s12884-024-06523-x>.

Supplementary Material 1

Acknowledgements

Not applicable.

Author contributions

The authors' contributions were as follows: Audêncio Victor and Patrícia H. Rondó: conceptualization, formal analysis, and methodology. Audêncio Victor: data curation. Patrícia H. Rondó: funding acquisition and supervision. Audêncio Victor, Liania A. Luzia, Leticia Falcão de Carvalho, Isabel Aires, Láisla Teles and Patrícia H. Rondó: investigation. Patrícia H. Rondó: project administration. Audêncio Victor and Patrícia H. Rondó: visualization and writing - original draft. Audêncio Victor, Liania A. Luzia, Leticia Falcão de Carvalho, Isabel Aires, and Patrícia H. Rondó: writing - review and editing. All authors contributed to the article and approved the submitted version.

Funding

This study was supported by the São Paulo Research Foundation (FAPESP) (grant number 2015/03333-6). AV, LFST, LFC and LDB had scholarships from the Coordination for the Improvement of Higher Education Personnel (CAPES). PPA had a scholarship from FAPESP (2018/17824-0).

Data availability

The datasets produced during this study will be accessible upon request directed to the corresponding author.

Declarations

Ethics approval and consent to participate

The research received ethical approval from the Research Ethics Committee with Human Subjects at the School of Public Health, University of São Paulo, before the commencement of data collection, as per protocol CAEE: 59787216.2.0000.5421 and opinion number 1.885.874. All participants provided informed consent, consistent with the principles outlined in the Helsinki Declaration.

Consent for publication

Not applicable.

Competing interests

The authors declare no competing interests.

Received: 1 January 2024 / Accepted: 16 April 2024

Published online: 25 April 2024

References

1. Champion ML, Harper LM. Gestational weight gain: update on outcomes and interventions. *Curr Diab Rep*. 2020;20:11.
2. Gesche J, Nilas L. Pregnancy outcome according to pre-pregnancy body mass index and gestational weight gain. *Int J Gynecol Obstet*. 2015;129:240–3.
3. O'Dwyer V, O'Toole F, Darcy S, Farah N, Kennelly MM, Turner MJ. Maternal obesity and gestational weight gain. *J Obstet Gynaecol (Lahore)*. 2013;33:671–4.
4. Kominiaiek MA, Lewkowitz AK, Carter E, Fowler SA, Simon M. Gestational weight gain and group prenatal care: a systematic review and meta-analysis. *BMC Pregnancy Childbirth*. 2019;19:18.
5. Kominiaiek MA, Peaceman AM. Gestational weight gain. *Am J Obstet Gynecol*. 2017;217:642–51.
6. IOM I of M. Nutrition during pregnancy: part I: Weight Gain, Part II: nutrient supplements. Washington, DC: The National Academies; 1990.
7. IOM. Weight Gain During Pregnancy: Reexamining the Guidelines, IOM. 2019;. (2019). Weight Gain During Pregnancy : Reexamining the Guidelines. Retrieved from www.iom.edu
8. Ren M, Li H, Cai W, Niu X, Ji W, Zhang Z, et al. Excessive gestational weight gain in accordance with the IOM criteria and the risk of hypertensive disorders of pregnancy: a meta-analysis. *BMC Pregnancy Childbirth*. 2018;18:281.
9. Macdonald-Wallis C, Tilling K, Fraser A, Nelson SM, Lawlor DA. Gestational weight gain as a risk factor for hypertensive disorders of pregnancy. *Am J Obstet Gynecol*. 2013;209:e3271–32717.
10. Whitaker KM, Ryan R, Becker C, Healy H. Gestational weight gain in twin pregnancies and maternal and Child Health: an updated systematic review. *J Womens Health (Larchmt)*. 2022;31:362–81.

11. Hasan SMT, Khan MA, Ahmed T. Inadequate maternal weight gain in the third trimester increases the risk of intrauterine growth restriction in rural Bangladesh. *PLoS ONE*. 2019;14:e0212116.
12. Wang L, Zhang X, Chen T, Tao J, Gao Y, Cai L, et al. Association of Gestational Weight Gain with Infant Morbidity and Mortality in the United States. *JAMA Netw Open*. 2021;4:e2141498.
13. Davis RR, Hofferth SL, Shenassa ED. Gestational weight gain and risk of infant death in the United States. *Am J Public Health*. 2014;104(Suppl 1):S90–5.
14. Truong YN, Yee LM, Caughey AB, Cheng YW. Weight gain in pregnancy: does the Institute of Medicine have it right? *Am J Obstet Gynecol*. 2015;212:362.e1–362.e8.
15. Cantor AG, Jungbauer RM, McDonagh M, Blazina I, Marshall NE, Weeks C, et al. Counseling and behavioral interventions for healthy weight and weight gain in pregnancy: evidence report and systematic review for the US Preventive Services Task Force. *JAMA*. 2021;325:2094–109.
16. Voerman E, Santos S, Patro Golab B, Amiano P, Ballester F, Barros H, et al. Maternal body mass index, gestational weight gain, and the risk of overweight and obesity across childhood: an individual participant data meta-analysis. *PLoS Med*. 2019;16:e1002744.
17. Aung W, Saw L, Sweet L. An integrative review of interventions for limiting gestational weight gain in pregnant women who are overweight or obese. *Women Birth*. 2022;35:108–26.
18. WHO WHO. Physical status: the use and interpretation of anthropometry. Geneva; 1995.
19. Stirnemann J, Villar J, Salomon LJ, Ohuma E, Ruyan P, Altman DG. International estimated fetal weight standards of the INTERGROWTH-21 st Project. 2017; March:478–86.
20. Villar J, Cheikh Ismail L, Victora CG, Ohuma EO, Bertino E, Altman DG, et al. International standards for newborn weight, length, and head circumference by gestational age and sex: the newborn cross-sectional study of the INTERGROWTH-21st Project. *Lancet*. 2014;384:857–68.
21. Textor J, Van der Zander B, Gilthorpe MS, Liškievicz M, Ellison GTH. Robust causal inference using directed acyclic graphs: the R package 'dagitty'. *Int J Epidemiol*. 2016;45:1887–94.
22. Shrier I, Platt RW. Reducing bias through directed acyclic graphs. *BMC Med Res Methodol*. 2008;8:1–15.
23. Hernán MA, Robins JM. Causal inference. 2010.
24. Ballinger GA. Using generalized estimating equations for longitudinal data analysis. *Organ Res Methods*. 2004;7:127–50.
25. Ceolin G, Moreira JD, Breda V, Mendes BC, Gomes FA, Mansur RB, et al. Association between adiposity and emergent depressive symptoms in a 10-years prospective cohort of older adults: the EpiFloripa Aging study. *J Affect Disord*. 2023;330:198–205.
26. Liang K-Y, Zeger SL. Longitudinal data analysis using generalized linear models. *Biometrika*. 1986;73:13–22.
27. Schubert J, Timmesfeld N, Noever K, Behnam S, Vinturache A, Arabin B. Impact of maternal body mass index and gestational weight gain on maternal and neonatal outcomes in twin pregnancies. *Acta Obstet Gynecol Scand*. 2023;102:181–9.
28. Goldstein RF, Abell SK, Ranasinha S, Misso M, Boyle JA, Black MH, et al. Association of Gestational Weight Gain with maternal and infant outcomes. *JAMA*. 2017;317:2207.
29. Langley-Evans SC, Pearce J, Ellis S. Overweight, obesity and excessive weight gain in pregnancy as risk factors for adverse pregnancy outcomes: a narrative review. *J Hum Nutr Diet*. 2022;35:250–64.
30. Liu X, Wang H, Yang L, Zhao M, Magnussen CG, Xi B. Associations between Gestational Weight Gain and adverse birth outcomes: a Population-based Retrospective Cohort Study of 9 million mother-infant pairs. *Front Nutr*. 2022;9.
31. Godoy AC, do Nascimento SL, Surita FG. A systematic review and meta-analysis of gestational weight gain recommendations and related outcomes in Brazil. *Clinics*. 2015;70:758–64.
32. Chiavaroli V, Hopkins SA, Biggs JB, Rodrigues RO, Seneviratne SN, Baldi JC, et al. The associations between maternal BMI and gestational weight gain and health outcomes in offspring at age 1 and 7 years. *Sci Rep*. 2021;11:20865.
33. Athukorala C, Rumbold AR, Willson KJ, Crowther CA. The risk of adverse pregnancy outcomes in women who are overweight or obese. *BMC Pregnancy Childbirth*. 2010;10:56.
34. Haugen M, Brantsæter AL, Winkvist A, Lissner L, Alexander J, Oftedal B, et al. Associations of pre-pregnancy body mass index and gestational weight gain with pregnancy outcome and postpartum weight retention: a prospective observational cohort study. *BMC Pregnancy Childbirth*. 2014;14:201.
35. Chowdhury R, N, Choudhary TS, Dhabhai N, Mittal P, Dewan R et al. Gestational weight gain and pregnancy outcomes: findings from north Indian pregnancy cohort. *Matern Child Nutr*. 2022;18.
36. Lipworth H, Barrett J, Murphy KE, Redelmeier D, Melamed N. Gestational weight gain in twin gestations and pregnancy outcomes: a systematic review and meta-analysis. *BJOG*. 2022;129:868–79.
37. Li H, Miao C, Xu L, Gao H, Bai M, Liu W, et al. Maternal pre-pregnancy body mass index, gestational weight gain trajectory, and risk of adverse perinatal outcomes. *Int J Gynecol Obstet*. 2022;157:723–32.
38. Montvignier Monnet A, Savoy D, Préaubert L, Hoffmann P, Bétry C. In underweight women, Insufficient Gestational Weight Gain is Associated with adverse obstetric outcomes. *Nutrients*. 2022;15:57.
39. Truong YN, Yee LM, Caughey AB, Cheng YW. Weight gain in pregnancy: does the Institute of Medicine have it right? *The American Journal of Obstetrics & Gynecology*. 2015;212:362.e1–362.e8.
40. Dodd J, Grivell R, Crowther C, Robinson J. Antenatal interventions for overweight or obese pregnant women: a systematic review of randomised trials. *BJOG*. 2010;117:1316–26.
41. Kac G, Velásquez-Meléndez G. Gestational weight gain and macrosomia in a cohort of mothers and their children. *J Pediatr (Rio J)*. 2005;81:47–53.
42. de Godoy AP, Viel CC, Melo EL, Dias DRC, Trevelin LC, Teixeira CAC. Multimedia Presentation integrating media with virtual 3D realistic environment produced in Real Time with High Performance Processing. *J Interact Syst*. 2014;5:1.
43. de Amorim MMR, Leite DFB, Gadelha TGN, Muniz AGV, Melo AS, de O. Rocha A Da M. Fatores De Risco para macrosomia em recém-nascidos de uma maternidade-escola no nordeste do Brasil. *Revista Brasileira De Ginecol E Obstetrícia*. 2009;31:241–8.
44. Godoy AC, Nascimento SL, Kasawara KT, Hatsue Oushiro N, Surita FG. A population-based study on gestational weight gain according to body mass index in the Southeast of Brazil. *Physiol J*. 2014;2014.
45. Padilha PDC, Accioly E, Chagas C, Portela E, Da Silva CL, Saunders C. Birth weight variation according to maternal characteristics and gestational weight gain in Brazilian women. *Nutr Hosp*. 2009;24:207–12.
46. Drehmer M, Duncan BB, Kac G, Schmidt MI. Association of Second and Third Trimester Weight Gain in pregnancy with maternal and fetal outcomes. *PLoS ONE*. 2013;8:e54704.
47. Nascimento SL do, Surita FG, Parpinelli de C, Cecatti MÂ, JG. Exercício físico no ganho de peso e resultados perinatais em gestantes com sobrepeso e obesidade: uma revisão sistemática de ensaios clínicos. *Cad Saude Publica*. 2011;27:407–16.
48. Hull HR, Thornton JC, Ji Y, Paley C, Rosenn B, Mathews P et al. Higher infant body fat with excessive gestational weight gain in overweight women. *Am J Obstet Gynecol*. 2011;205:211.e1–211.e7.
49. Godfrey KM, Reynolds RM, Prescott SL, Nyirenda M, Jaddoe VWW, Eriksson JG, et al. Influence of maternal obesity on the long-term health of offspring. *Lancet Diabetes Endocrinol*. 2017;5:53–64.
50. Desai M, Jellyman JK, Ross MG. Epigenomics, gestational programming and risk of metabolic syndrome. *Int J Obes*. 2015;39:633–41.
51. Starling AP, Brinton JT, Glueck DH, Shapiro AL, Harrod CS, Lynch AM, et al. Associations of maternal BMI and gestational weight gain with neonatal adiposity in the healthy start study. *Am J Clin Nutr*. 2015;101:302–9.
52. Mina TH, Lahti M, Drake AJ, Räikkönen K, Minnis H, Denison FC, et al. Prenatal exposure to very severe maternal obesity is associated with adverse neuropsychiatric outcomes in children. *Psychol Med*. 2017;47:353–62.
53. Catalano P, Ehrenberg H. Review article: the short- and long-term implications of maternal obesity on the mother and her offspring. *BJOG*. 2006;113:1126–33.

Publisher's Note

Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.